éggagagcc ccggagtgag cggagtagcg agtcggcaac ccggaggggt agaaatattt 60 ctgtc atg gct cat tca aag act agg acc aat gat gga aaa att aca tat 110 Met Ala His Ser Lys Thr Arg Thr Asn Asp Gly Lys Ile Thr Tyr ccg cct ggg gtc aag gaa ata tca gat aaa ata tct aaa gag gag atg 158 Pro Pro Gly Val Lys Glu Ile Ser Asp Lys Ile Ser Lys Glu Glu Met 20 -25 30 206 gtg aga cga tta aag atg gtt gtg aaa act ttt atg gat atg gac cag Val Arg Arg Leu Lys Met Val Val Lys Thr Phe Met Asp Met Asp Gln 35 40 gac tot gaa gaa gaa aag gag ott tat tta aac ota got tta cat ott 254 Asp Ser Glu Glu Glu Lys Glu Leu Tyr Leu Asn Leu Ala Leu His Leu 50 get tea gat ttt ttt etc aag cat eet ggt aaa gat gtt ege tta etg 302 Ala Ser Asp Phe Phe Leu Lys His Pro Gly Lys Asp Val Arg Leu Leu 65 70 gta gcc tgc tgc ctt gct gat att ttc agg att tat gct cct gaa gct 350 Val Ala Cys Cys Leu Ala Asp Ile Phe Arg Ile Tyr Ala Pro Glu Ala 80 85 cct tac aca tcc cct gat aaa cta aag gat ata ttt atg ttt ata aca 398 Pro Tyr Thr Ser Pro Asp Lys Leu Lys Asp Ile Phe Met Phe Ile Thr 100 105 110 446 aga cag ttg aag ggg cta gag gat aca aag agc cca caa ttc aat agg Arg Gln Leu Lys Gly Leu Glu Asp Thr Lys Ser Pro Gln Phe Asn Arg 115 120 125 tat ttt tat tta ctt gag aac att gct tgg gtc aag tca tat aac ata 494 Tyr Phe Tyr Leu Leu Glu Asn Ile Ala Trp Val Lys Ser Tyr Asn Ile 130 140 135 542 tgc ttt gag tta gaa gat agc aat gaa att ttc acc cag cta tac aga Cys Phe Glu Leu Glu Asp Ser Asn Glu Ile Phe Thr Gln Leu Tyr Arg 155 145 150

DEC 0 1. 2005

acc Thr 160	tta Leu	ttt Phe	tca Ser	gtt Val	ata Ile 165	aac Asn	aat Asn	ggc Gly	cac His	aat Asn 170	cag Gln	aaa Lys	gtc Val	cat His	atg Met 175	590
cac His	atg Met	gta Val	gac Asp	ctt Leu 180	atg Met	agc Ser	tct Ser	att Ile	att Ile 185	tgt Cys	gaa Glu	ggt Gly	gat Asp	aca Thr 190	gtg Val	638
tct Ser	cag Gln	gag Glu	ctt Leu 195	ttg Leu	gat Asp	acg Thr	gtt Val	tta Leu 200	gta Val	aat Asn	ctg Leu	gta Val	cct Pro 205	Ala	cat His	686
aag Lys	aat Asn	tta Leu 210	aac Asn	aag Lys	caa Gln	gca Ala	tat Tyr 215	gat Asp	ttg Leu	gca Ala	aag Lys	gct Ala 220	tta Leu	ctg Leu	aag Lys	734
agg Arg	aca Thr 225	gct Ala	caa Gln	gct Ala	att Ile	gag Glu 230	cca Pro	tat Tyr	att Ile	acc Thr	act Thr 235	ttt Phe	ttt Phe	aat Asn	cag G1n	782
gtt Val 240	ctg Leu	atg Met	ctt Leu	999 Gly	aaa Lys 245	aca Thr	tct Ser	atc Ile	agc Ser	gat Asp 250	ttg Leu	tca Ser	gag Glu	cat His	gtc Val 255	830
ttt Phe	gac Asp	tta Leu	att Ile	ttg Leu 260	gag Glu	ctc Leu	tac Tyr	aat Asn	att Ile 265	gat Asp	agt Ser	cat His	ttg Leu	ctg Leu 270	ctc Leu	878
tct Ser	gtt Val	tta Leu	ccc Pro 275	cag Gln	ctt Leu	gaa Glu	ttt Phe	aaa Lys 280	Leu	aag Lys	agc Ser	aat Asn	gat Asp 285	Asn	gag Glu	926
gag Glu	cgc Arg	cta Leu 290	Gln	gtt Val	gtt Val	aaa Lys	cta Leu 295	Leu	gca Ala	aaa Lys	atg Met	ttt Phe 300	Gly	gca Ala	aag Lys	974
gat Asp	tca Ser 305	Glu	ttg Leu	gct Ala	tct Ser	caa Gln 310	Asn	aag Lys	cca Pro	ctt Leu	tgg Trp 315	Gln	tgc Cys	tac Tyr	ttg Leu	1022
990 Gly 320	Arg	ttt Phe	aat Asr	gat Asp	atc 11e 325	His	gta Val	cca Pro	ato Ile	cgc Arg 330	Leu	gaa Glu	tgt Cys	gtg Val	aaa Lys 335	1070

	-	_		-		atg Met		His		_		_		-	Leu	1118
						agg Arg			_			_	-		-	1166
	-	_		_		ata Ile	-		_	-		_	-			1214
Leu	_		-			ctt Leu 390				-		_			_	1262
				Val		aaa Lys										1310
	_			_		cag Gln		-	-			_	_			1358
_						gac Asp		_								1406
-				_		ctt Leu	_	_				_			_	1454
						act Thr 470		-		_		-				1502
				-	-	tta Leu					-	-		-		1550
_		_			_	ctc Leu	_			-	_	_	_		_	1598

		aag Lys												1646
		atg Met 530				Leu								1694
		atg Met												1742
		cag Gln												1790
		ggt Gly												1838
		aca Thr												1886
	Ala	cct Pro 610												1934
		aac Asn												1982
_		act Thr	-	_	_				_	_		_	gta Val 655	2030
		ttt Phe												2078
		ctg Leu		_	_		_	_		-	_	-	-	2126

		caa Gln 690												2174
		atc Ile												2222
		ccc Pro	-		-			_			_			2270
		tct Ser					Phe							2318
	_	agc Ser	_		-		_	-					_	2366
-		att Ile 770			_			-		_			 _	2414
		aaa Lys												2462
	_	cgg Arg				-						-	-	2510
		gta Val		-		_	-		Пe	-	_			2558
		cga Arg	Leu								-			2606
		acc Thr 850	_	_					_		_	-		2654

	_		_					-	aaa Lys								2702
•		-	-			-			aag Lys	_		Gln					2750
		-					-		tat Tyr	_		-	-		_	Ile	2798
		-	Glu				_	_	caa Gln 920			_	_			cac His	2846
					-				cca Pro				-			-	2894
	_		_	•		_		-	aag Lys			_			-		2942
		_	_						gta Val			Glu		_	_	cag Gln 975	2990
		-	-	_	. –	_			ttg Leu							_	3038
								Leu	gca Ala 1000				Asp			aaa Lys	3086
		Gln	-				Leu		gat Asp			Glu	-			ttt Phe	3134
	Val	_	-			Met	_		aat Asn	_	Asn		_		-		3182

	Arg	_	atg Met	Val	-				Gln			-		Gln		3230
cca Pro	_	_	Āla		_		_	Lys	-			_	Cys			3278
gcc Ala	_	Asn			•		Lys	•				Seri	-	_		3326
cct Pro	Lys					Pro					Thr					3374
aat Asn 1		-			Lys		Tyr	_		Pro	-	_				3422
ttc Phe 1120	Thr		gga Gly	Lys					Asn	-				Val		3470
			tca Ser		Ala			Gln					Ser			3518
_	_	Thr	gta Val 1155			-	Ser					Pro				3566
	Arg		aag Lys			Leu					Met				gaa Glu	3614
Asn			tac Tyr		Met					Pro						3662
	Arg		gac Asp	Ser					Ser		Leu			Pro		3710

FIG. I-7

•	ggc Gly	agg Arg	aaa Lys	Lys	acg Thr 220	Pro	gtc Val	aca. Thr	Glu	cag Gln 225	gag Glu	gag Glu	aaa Lys	Leu	ggt Gly 1230	atg Met	3758
	gat Asp	gac Asp	Leu	act Thr 1235	aag Lys	ttg Leu	gta Val	Gln	gaa Glu 240	cag Gln	aaa Lys	cct Pro	Lys	ggc Gly L245	agt Ser	cag Gln	3806
	cga Arg	Ser	cgg Arg L250	aaa Lys	aga Arg	ggc Gly	His	acg Thr 1255	gct Ala	tca Ser	gaa Glu	tct Ser	gat Asp 1260	gaa Glu	cag Gln	cag Gln	3854
	Trp	cct Pro .265	gag Glu	gaa Glu	aag Lys	Arg	ctc Leu 270	aaa Lys	gaa Glu	gat Asp	Ile	tta Leu L275	gaa Glu	aat Asn	gaa Glu	gat Asp	3902
		Gln			Pro					Lys		ggc Gly			Pro		3950
	cct Pro	ctt Leu	ggt Gly	Gly	ggt Gly L300	aca Thr	cca Pro	aaa Lys	Glu	gag G1u L305	cca Pro	aca Thr	atg Met	Lys	act Thr 1310	tct Ser	3998
			Gly		Lys			Ser				gca Ala	Pro				4046
		Glu					Ser					cag Gln					4094
	Lys	cag Gln 1345	cac His	cga Arg	gtg Val	Ser	agg Arg 1350	aga Arg	gca Ala	cag Gln	Gln	aga Arg 1355	gca Ala	gaa Glu	tct Ser	cct. Pro	4142
		Ser			Ile					Ser		cca Pro			Gly		4190
	gga Gly	aga Arg	cca Pro	Ser	aaa Lys 1380	Thr	cca Pro	tca Ser	Pro	tca Ser 1385	Gln	cca Pro	aaa Lys	Lys	aat Asn 1390	Val	4238

taagttgtaa atattacatt tcaaaccaat ttcaaattat tttgcaaaag ttcctaaatt 4298 End

tgtaaacata catattgctg tatttaaatt ccatatattt agccccatta cactaggtac 4358 ggcggcgaag tgctaaaagg gaacggcgat gaacaaatgt aattaataac tttctctgtg 4418 aaagctttgg aaaaatcttt ttttttttt ttttttttg gtcaagcttg aggctgaata 4478 aagcctttga tgcacaaaat gggactgctg aagagtggac agttggacct tactttggtg 4538 accccataca tttgtggtca catgctttag ccatacacat ggtaacattg actatggagt 4598 cttgtgaaag tgtaatgtgc gatggctatg tagacataaa gaagaaactt gtaaatatct 4658 tttttctttt ttttaatqtt tctqatttct gaagtgcttg tatagctttt atctgcggct 4718 ttaaactgac agtacccgac tgtttattgg atctattgat ttgaaaagaa tttgttagga 4778 tagatettaa geagtaatet geegtgetti geattigeat titetge<u>aat tit</u>actgiga 4838 aaaaaaattt gttttcaaca attggtgtca ttttcttgat gtcactattt gttggagagt 4898 taaatggtet etteeettig tgtatettae etagtgttta eteetgggea eeettaatet 4958 tcagaggtgc taaattgtct gccattacac cagaaggatg cctctgatag gaggacaacc 5018 atgcaaattg tgaaatagtc ctgaagttct tggattactt tacacctcag tattgatttg 5078 tcccagaatt ttctggcctt tcatggcaat gaaaatttta agaagaaaga tttaaagtat 5138 tttaatttta aagagtgtgt tataaaataa tgtactgaat tctttatccc attttatcat 5198 cctttcagtt tttattaatc tactgtatca ataaaattct gtaatttgaa tgagtaaaaa 5258 5271 aaaaaaaaa aaa



319 196 LDTVLVNLVPAHKNLNKQAYDL LYLNLALHLASDFFLKHPGKDVRLLVACCLADIFRIYAPEAPYTSPDKLKDIFMFITRQLKGL 217 LMLGKTS ISDLSEHVFDLILELYNIDSHLLLSVLPQL 355 375

277

LGRFNDIHVPIRLECVKFASHCLMNHPDLAKDLTEYL

VTAAKKDILLVNDHLLNFVRERTLDKRWRV

Similar protein kinase sequences:	Consensus: β-strand AS3 position: 419 AS3 sequence: YALQ.SA	conserved regions:
YTLGVSA (Elm 1) YALLNLL (Tsl) YHLKQNI (Cdc15) YKLVRKI (CK1a IVLQESI (Alk5)	β-strand 419 YALQ.SA	
YTIGVSA (Elm 1) GeDrfGkV (Ror2) YALLNLL (Tsl) GsGsfGdI (CK1a) YHLKQNI (Cdc15) AeGesHiS (Ypka) YKLVRKI (CK1a HeSdfSeV (Mik1) IVLQESI (Alk5)	Mg-ATP binding loop β-strand 2 (G x G x x G x V) 426 453 GKDAAKQI LLVERIE	Subdomain I
LLYELMD (Yk1516) YLGEQVS (PKN2) YLCLCLN (BCK1)	β-strand 2 453 LLVERIF	
LLYELMD (Yk1516) YAM K CLKKDVI (CeTPAI) QAFKN K MQVL (Araf) YLGEQVS (PKN2) YAM K CLDKKRI (bARKI) TLALN E RIML (bARKI) YLCLCLN (BCK1) YAM K CLDKKRI (DmGPRKI) YTRVR E IKFI (SMEI) VAI K CIAKKAL (CamKI)	β-strand 3 (x x x K x x x) 472 ERM K CLYYLYA	Subdomain II
QAFKN K MQVL (Araf) TLALN E RIML (bARKI) YTRVR E IKFI (SMEI)	α-helix C (x x x x E x x x) 489 VKALN E MWKC	Subdomain III

Hank's

Hank's conserved

AS3 sequence:	AS3 position:	Consensus:
LIDIVKDP	509	β-strand 4
IFSK.VM	525	β-strand 5

70
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regions:

Similar protein kinase sequences:	AS3 position: AS3 sequence:	Consensus:
LLDIVKDP (TPCKII) LLDWFERP (Pim1) LLGLCREA (Klg) LVKLIGYC (APK1)	509 LIDIVKDP	β-strand 4
IFSCLVME (PvpK1) KFSCLVME (G11a) KFSCLVME (ZmPP)	525 IFSK.VMV	β-strand 5

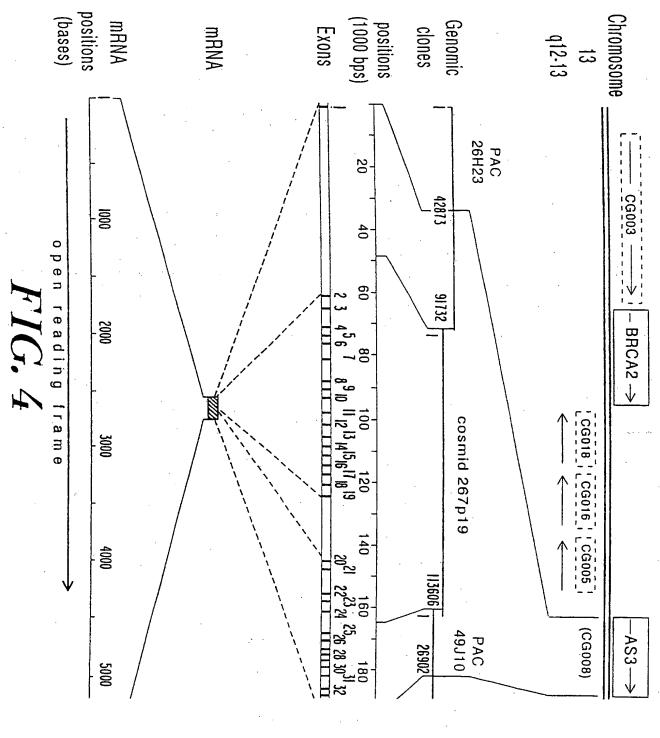
Subdomain V

Subdomain VIa

525 IFSK.VMV	β-strand 5
540 GKAQDEMKK	α-helix D
554 EDDEKIRKQ LEVL	α-helix E

PK) GNLQEYLTR (TGFbRII) GNLQNFLKL (Let23)
GSLQNFLRE (TORSO) ERDADAVKQILEA (CaMKIV)
ECDANIMKQILSG (PfCPK)
ADQLNIAKQISAG (TORRTK)

ESVIMYTKQLLL (NPK1)



{42873}	1 CCGGAGAG	Exon 1ACCCGG	46 {42919} AG * gtaggaa
(13347) ttttcttgtttcag *			73 (13475) AG * gtgagta
(16397) ttttatttttgtatag *	174 ATGGTTGT		77 (16602) AG * gcaagta
(22832) tcttttttttatttaag *	378 GATATATT		64 (22920) AG * gtaagca
(23028) ccttatttttag *	465 AACATTGC	5 Exon 5GTTATA	52 (23125) AA * gtaagtt
(23747) ttttgaattgcag *		Exon 6CTCATA	39 (23873) AG * gtgagta
(32357) tttatgtttttcag *	690 AATTTAAA	Exon 7TTACCA	54 (32439) CT * gtaagtc
(37809) ctttctcctcaaaag *			11 (37951) AG * gtaactt
(40437) ttttattttag *	912 AGCAATGA		27 (40554) AG * gtatatg
(43428) tttatattttatcag *			22 (43524) AG * gtactat
(48471) tgttatctttcag *			58 (48617) GA * gtaagta
(51727) ttttttgtttttaag *			20 (51880) CG * gtaagtt
(53049) tctgcttttttgtag *			34 (53164) WA * gtatgtt

FIG. 5-1

(58816)tttgtgtttttcag '	1535 AGCATTGA	Exor	14	1616 AACCCAAA	(58898) * gtaagta
(61447) ttgtgtgatttacag *	1617 ACAGATGC	Exon	15	1665 TATTACAA	(61497) * gtaagtt
(64323) tttattttaag *	1666 GAAATTTA	Exon	16	1805 GTTGTGTG	(64464) * gtaagga
(65916) taatctgtattacag *	1806 CGTGAAAT	Exon	17	1921 TCTATCAG	(66033) * gtatttg
(71527) ttggtcatattttag *	1922 TGCTCTTA	Exon	18	2027 TGCTTAAG	(71633) * gtaagta
(74539) tgattcattttatag *		Exon	19	2188 ATCAGATC	(74700) * gtgagtt
(96694) tttttttttaatag *	2189 AGCCTTGC	Exon	20	2312 TATTTGAG	(96818) * gtaatga
(99765) tcccctcattttcag *	2313 CCTCTGCA	Exon	21	2471 ATGATCGG	(99925) * gtaattt
(105674) ctcgtttatttttag *	2472 CTTCCAGG	Exon	22	2540 TGGTCAAA	(105744) * gtgagta
(107185) ttgtctcttaaatag *				2677 AAAATTAG	(107322) * gtatgca
(110571) ctactcatttttcag *		Exon	24	2801 CTATCAAC	(110696) * gtaagga
[4319] ttgtgtctttacag *	2802 GATGAATG	Exon	25	3006 TGTTAGTG	[4524] * gtaagca

FIG. 5-2

[6829] ttttctttttcag *			3121 26GTTAAAGA	
[9074] ttttttttttttag *	3122 ATGTCTTT	Exon	3254 27TGAATGAA	[9208] * gtatgta
[9522] tatactattgcag *	3255 AAACTGTA			[9642] * gtagtta
[10614] ttctcttggttgtag *	3375 AATTTCAG	Exon	3437 29CTGGAAAA	[10679 * gtatgtt
[11561] catttctcatttcag *	3438 CCTAAAAC	Exon		[11709] * gtaagtg
[15476] tgtctgtattaaaag *	3584 GCTTGATA			[15583] * gtgagat
_21107] tttttttttcccctag *	3690 TCTGAATT	Exon		[21548] * gtaagca
[21640] tcttccccaaagcag *	4130 AGCAGAAT			[21866] * gtaagat
[26002]ctttccttttaag *		Exon		[26902] * (poly-A)

FIG. 5-3